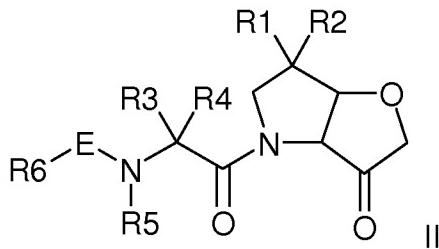


Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application.

1. (Currently amended) A compound of the formula II



wherein

one of R¹ and R² is halo and the other is H or halo;

R³ is C₁-C₅ straight or branched chain, optionally fluorinated, alkyl;

R⁴ is H; or

R³ together with R⁴ defines

a spiro-C₅-C₇ cycloalkyl, optionally substituted with 1 to 3 substituents selected from halo, hydroxyl, C₁-C₄ alkyl or C₁-C₄ haloalkyl; or optionally bridged with a methylene group; or

a C₄-C₆ saturated heterocycle having a hetero atom selected from

O, NR_a, S, S(=O)₂ ;

R⁵ is independently selected from H or methyl;

E is -C(=O)-, -S(=O)_m-, -NR⁵S(=O)_m-, -NR⁵C(=O)-, -OC(=O)-,

R⁶ is a stable, optionally substituted, monocyclic or bicyclic, carbocycle or heterocycle

heterocycle wherein the or each ring has 4, 5 or 6 ring atoms and 0 to 3 hetero atoms selected

from S, O and N and wherein the optional substituents comprise 1 to 3 members selected from

R₇ R⁷:

R₇ R⁷ is independently selected from halo, oxo, nitrile, nitro, C₁-C₄ alkyl, [[-XNRaRb,]] -X-NRbR⁹, [[-NRb-XNRaRb-R⁹]] -NRb-X'-R⁹, NH₂CO-, -X-R⁹, -X-O-R⁹, [[O-X-R⁹]] -O-X'-R⁹, -X-C(=O)R⁹, -X-(C=O)NRaR⁹, -X-NRbC(=O)R⁹, -X-NHSO_mR⁹, -X-S(=O)_mR⁹, -X-C(=O)OR⁹, -X-NRbC(=O)OR⁹;

R₉ R⁹ is independently H, C₁-C₄ alkyl, C₃-C₆ cycloalkyl, pyrrolidinyl, piperidinyl, morpholinyl, thiomorpholinyl, piperazinyl, indolinyl, pyranyl, thiopyran, furanyl, thienyl, pyrrolyl, oxazolyl, isoxazolyl, thiazolyl, imidazolyl, pyridinyl, pyrimidinyl, pyrazinyl, indolyl, phenyl, any of which is optionally substituted with R¹⁰;

R₁₀ R¹⁰ is independently selected from hydroxy, ~~X~~R⁹, ~~XNRaRb~~, ~~XNRbR⁹~~, ~~NRbC₄-~~ C₄alkylR⁹ -X-R⁹, -X-NRbR⁹, -NRb-X'-R⁹, nitro, cyano, carboxy, oxo, [[C₁-C₄ alkyl,]] C₁-C₄-alkoxy, C₁-C₄ alkanoyl, carbamoyl;

R⁹ is independently H, C₁-C₄ alkyl, C₃-C₆ cycloalkyl, pyrrolidinyl, piperidinyl, morpholinyl, thiomorpholinyl, piperazinyl, indolinyl, pyranyl, thiopyran, furanyl, thienyl, pyrrolyl, oxazolyl, isoxazolyl, thiazolyl, imidazolyl, pyridinyl, pyrimidinyl, pyrazinyl, indolyl, phenyl, any of which is optionally substituted with R¹⁰;

R¹⁰, is independently selected from hydroxy, nitro, cyano, carboxy, oxo, C₁-C₄ alkyl, C₁-C₄-alkoxy, C₁-C₄ alkanoyl, carbamoyl; or, where R⁶ is a monocyclic group substituted directly or via methylene by an aryl or a 5 or 6 membered heteroaryl moiety substituted by R⁹, which is a morpholinyl, piperidinyl or piperazinyl group, then R¹⁰, can additionally be fluoro, difluoro, or C₁-C₃alkyloxyC₁-C₃alkyl-; or, where R⁶ is phenyl substituted by thiazol-4-yl, 5-methylthiazol-4-

yl or thien-2-yl, any of which is substituted by morpholinylmethyl-, piperidinylmethyl-, piperazinylmethyl-, then R¹⁰, may additionally be fluoro, difluoro or C₁-C₃ alkyl-O-C₁-C₃alkyl-

X is independently a bond or C₁-C₄ alkyl alkylenyl;

X' is independently C₁-C₄ alkylenel;

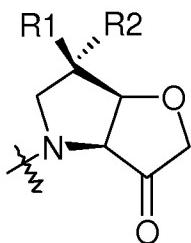
Ra is independently H, C₁-C₄ alkyl or CH₃C(=O);

Rb is independently H, or C₁-C₄ alkyl

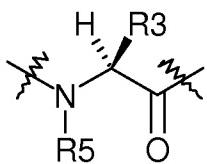
m is independently 0,1 or 2;

or a pharmaceutically acceptable salt or prodrug solvate thereof.

2. (Original) A compound according to claim 1, wherein the stereochemistry is as depicted in the partial structure below:



3. (Original) A compound according to claim 1, wherein the stereochemistry is as depicted in the partial structure below:



4. (Original) A compound according to claim 1, wherein R² is halo and R¹ is H.

5. (Original) A compound according to claim 4, wherein R² is fluoro.

6. (Original) A compound according to claim 1, wherein R¹ and R² are fluoro.

7. (Original) A compound according to claim 1, wherein R³ is C₁-C₄ branched chain alkyl.

8. (Original) A compound according to claim 7, wherein R³ is iso-butyl.

9. (Original) A compound according to claim 1, wherein R³ and R⁴ together define spirocycloalkyl.

10. (Original) A compound according to claim 9, wherein R³ and R⁴ together define spirocyclohexyl.

11. (Original) A compound according to claim 1, wherein R⁵ is H.

12. (Original) A compound according to claim 1, wherein E is -C(=O)-.

13. (Original) A compound according to claim 1, wherein R⁶ is substituted phenyl.

14. (Original) A compound according to claim 13, wherein the substituent comprises -NRaRb, -CH₂NRaRb, -NRbR⁹, -NRbC₁-C₄alkylR⁹, C₁-C₄ straight or branched alkyl or -O-R⁹.

15. (Original) A compound according to claim 14, wherein the substituent comprises -NH-CH₂phenyl, -NHCH₂pyridyl or -NH-phenyl, wherein each phenyl or pyridyl ring is substituted with C₁-C₄-alkyl, -NRaRb, -NRbR⁹ or -NRbC₁-C₄alkylR⁹.

16. (Original) A compound according to claim 13, wherein the substituent comprises C₃-C₆ cycloalkyl, pyrrolidinyl, piperidinyl, morpholinyl, thiomorpholinyl, piperazinyl, indolinyl, pyranyl, thiopyranyl, furanyl, thienyl, pyrrolyl, oxazolyl, isoxazolyl, thiazolyl, imidazolyl, pyridinyl, pyrimidinyl, pyrazinyl, indolyl, phenyl, any of which is optionally substituted with R¹⁰.

17. (Original) A compound according to claim 16, wherein the substituent is selected from indolinyl, pyranyl, thiopyranyl, pyrrolyl, oxazolyl, isoxazolyl, thiazolyl, imidazolyl, pyridinyl, pyrimidinyl, pyrazinyl, indolyl, any of which is optionally substituted with R¹⁰.

18. (Original) A compound according to claim 17, wherein the substituent is thiazolyl, 5-methyl-thiazolyl or thienyl, optionally substituted with R¹⁰.

19. (Original) A compound according to claim 18, wherein the substituent is thiazol-4-yl, 5-methylthiazol-4-yl or thien-2-yl, optionally substituted with R¹⁰.

20. (Original) A compound according to claim 18, wherein the thiazolyl, 5-methylthiazolyl or theinyl is substituted with morpholinyl, morpholinylmethyl-, piperidinyl, piperidinylmethyl-, piperazinyl, piperazinylmethyl, any of which is substituted with C₁-C₃ alkyl, fluoro, difluoro or C₁-C₃ alkyl-O-C₁-C₃alkyl-.

21. (Original) A compound according to claim 20, wherein the substituent to the thiazolyl, 5-methylthiazolyl or thienyl is piperid-4-yl which is substituted with methyl, piperazinyl which is N-substituted with C₁-C₃ alkyl or methyloxyethyl-, -or piperid-1-ylmethyl- which is unsubstituted or 4-substituted with fluoro or di-fluoro.

22. (Original) A compound according to claim 13, wherein the substituent comprises a morpholine, piperidine or piperazine ring, optionally substituted with R¹⁰.

23. (Original) A compound according to claim 22 comprising piperid-4-yl or N-piperazinyl, N-substituted with Ra or piperidin-1-yl which is 4-substituted with -NRaRb.

24. (Original) A compound according to claim 1, wherein R⁶ is optionally substituted: benzothiazol or benzofuryl or benzoxazolyl.

25. (Currently amended) A compound according to claim 24, wherein the substituent is -OR⁹, [[-OXR⁹]] -O-X'-R⁹, -NRbR⁹ or [[-NRbXR⁹]] -NRb-X'-R⁹.

26. (Original) A compound according to claim 25, wherein R⁹ is piperid-4-yl, piperazin-1-yl or piperidin-1-yl or morpholino, any of which is substituted with C₁-C₃ alkyl.

27. (Original) A compound according to claim 26, wherein the optional substituent to R⁶ is N-morpholinylethoxy, N-methylpiperid-4-yloxy, or N-methylmorpholin-3-ylmethoxy.

28. (Previously presented) A pharmaceutical composition comprising a compound as defined in claim 1 and a pharmaceutically acceptable carrier or diluents therefor.

29. (Withdrawn) A method for the treatment of a disorder mediated by cathepsin K comprising administering a compound as defined in claim 1.

30. (Withdrawn) A method according to claim 29, wherein the disorder is selected from:

osteoporosis,

gingival diseases such as gingivitis and periodontitis,

Paget's disease,

hypercalcaemia of malignancy

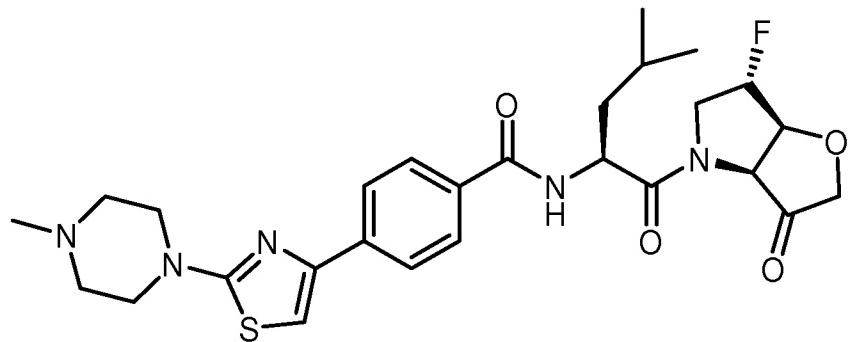
metabolic bone disease

diseases characterised by excessive cartilage or matrix degradation, such as osteoarthritis and rheumatoid arthritis.

bone cancers including neoplasia,

pain.

31. (New) A compound according to claim 1 which is:



or a pharmaceutically acceptable salt or solvate thereof.